

## Supplemental Online Content

Coburn SB, Humes E, Lang R, et al; Corona-Infectious-Virus Epidemiology Team (CIVETs) of the NA-ACCORD of IeDEA. Analysis of postvaccination breakthrough COVID-19 infections among adults with HIV in the United States. *JAMA Netw Open*. 2022;5(6):e2215934. doi:10.1001/jamanetworkopen.2022.15934

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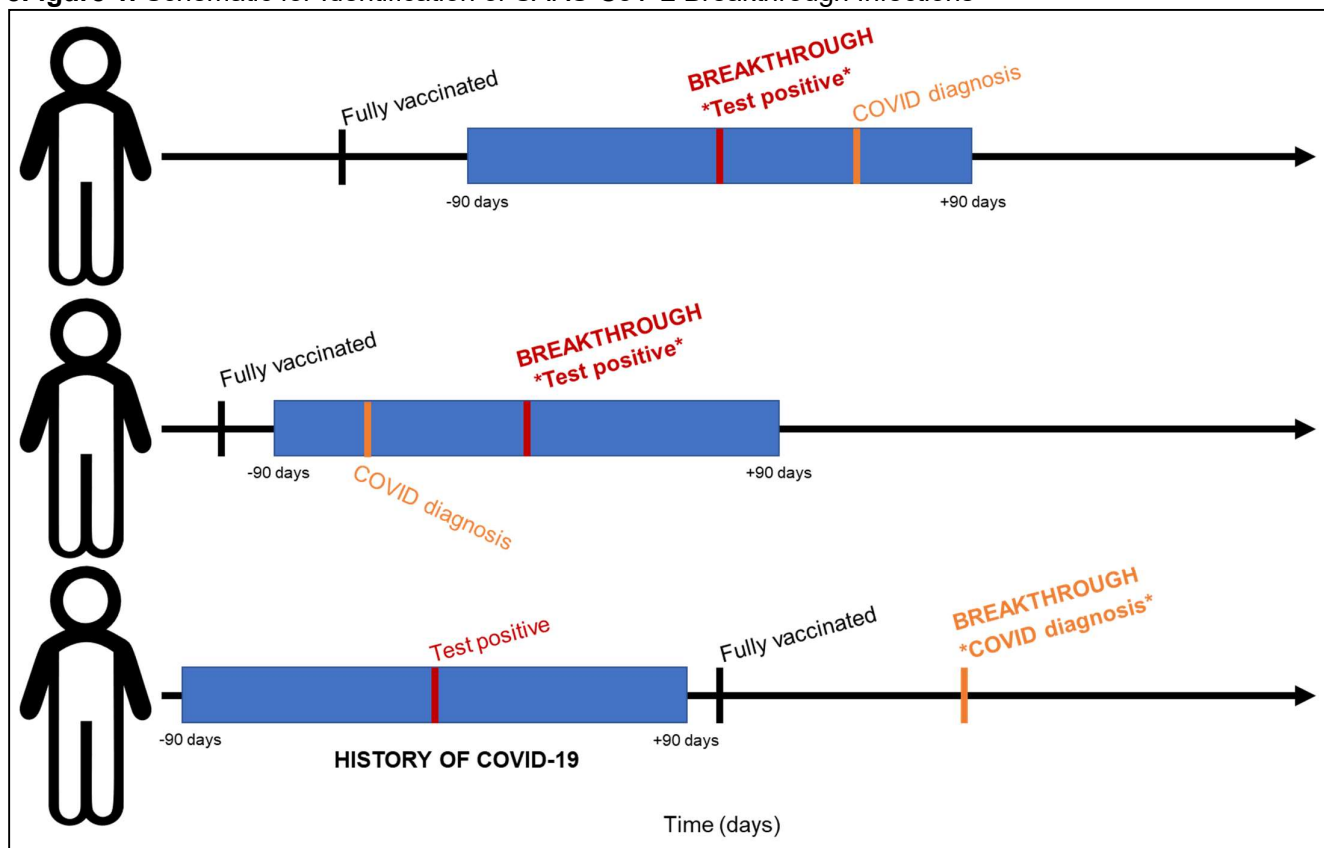
This supplemental material has been provided by the authors to give readers additional information about their work.

**eTable 1.** Definitions of in Care and Criteria Used to Identify People With and Without HIV

| <b>Cohort</b>                                  | <b>Description</b>   | <b>“In care” or “in cohort”</b>                                    | <b>PWH</b>  |
|--|--|--|---|
| Kaiser Permanente Mid-Atlantic States (KPMAS)  | Integrated Health system   | In Care – KPMAS membership for ≥1 month in 2020                    | HIV registry*   |
| Kaiser Permanente Northern California (KPNC)   | Integrated Health system   | In Care – KPNC membership for ≥1 month between 3/1/20 and 12/31/20 | HIV registry*   |
| University of North Carolina Chapel Hill (UNC) | Medical center cohort  | ≥1 encounter with UNC Health in 2019 and alive as of 03/01/2020    | HIV diagnosis (by ICD diagnosis code)                                     |
| Veterans Aging Cohort Study (VACS)             | National cohort sampling from all PWH and 1:2 demographically-matched PWoH in care in the Veterans Health Administration (VA) system | Enrolled in VACS from 1996-2017 and alive in 2020                  | HIV diagnosis (presence of 1 inpatient or 2 outpatient ICD codes for HIV) |

\*The Kaiser Permanente Mid-Atlantic States and Northern California HIV Registries are databases of members diagnosed with HIV since 1998 and 1980, respectively. Primary sources used to identify HIV patients are HIV-specific laboratory tests, antiretroviral therapy, hospital-based HIV diagnosis, and diagnosis confirmed by infectious disease physicians.

**eFigure 1.** Schematic for Identification of SARS-CoV-2 Breakthrough Infections



This figure depicts three mock scenarios to demonstrate how we defined breakthrough infection using laboratory data and diagnosis codes. Each “person” demonstrates possible scenarios of when and how laboratory tests were confirmed/diagnoses were made, and how we reconciled these records. For each patient, all positive SARS-CoV-2 laboratory tests were identified. Additional positive laboratory tests and/or diagnoses occurring within +/- 90 days of the date of a positive/detectable SARS-CoV-2 test result were considered persistent infection per CDC suggested classification. In instances where there was a COVID-19 diagnosis within the 90-day window of a positive or detectable test result, the date of the laboratory test was used to define the date of the breakthrough case; positive laboratory results were prioritized over diagnosis codes for calculation of breakthrough date due to their greater specificity. This reconciliation was repeated for diagnoses that did not occur within +/- 90 window of another laboratory test.

**Footnotes:**

Blue bar indicates +/- 90 window around a positive or detectable SARS-CoV-2 test result.

Red bar/text indicates the date of a positive or detectable SARS-CoV-2 test result.

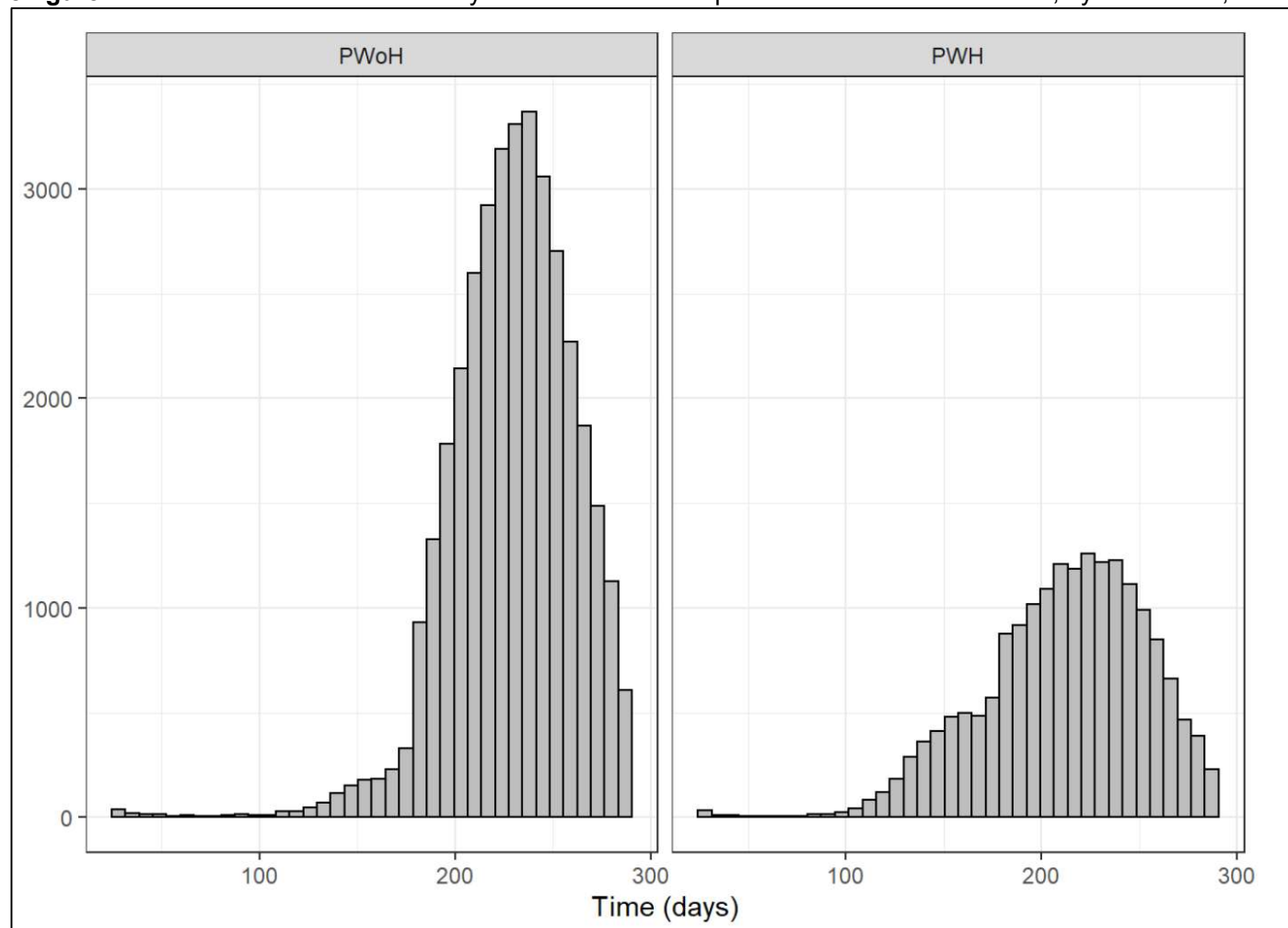
Orange bar/text indicates the date of a COVID-19 diagnosis (using ICD-10 codes).

Diagnosis codes include: U07.1 (specific to COVID-19), B34.2 (Coronavirus infection, unspecified), B97.21 (SARS-associated coronavirus causing disease classified elsewhere), B97.29 (other coronavirus as the cause of diseases classified elsewhere, or J12.81 (pneumonia due to SARS-associated coronavirus).

**eTable 2.** Timeline of Vaccination Availability and Vaccine Guidelines Recommendations, December 2020 to December 2021

| Date                               | Action  |
|------------------------------------|---|
| <a href="#">December 11, 2020</a>  | FDA approves the Pfizer mRNA COVID-19 vaccine   |
| <a href="#">December 18, 2021</a>  | FDA approves Moderna mRNA COVID-19 vaccine  |
| January 18, 2021                   | First date Pfizer recipients could be “fully vaccinated” (initial dose, second dose 21 days later, 14 days after 2 <sup>nd</sup> dose)  |
| January 22, 2021                   | First date Moderna recipients could be “fully vaccinated” (initial dose, second dose 28 days later, 14 days after 2 <sup>nd</sup> dose)   |
| <a href="#">February 27, 2021</a>  | FDA approves J&J COVID-19 vaccine   |
| March 13, 2021                     | First date J&J recipients could be “fully vaccinated” (14 days after 1 <sup>st</sup> dose)  |
| <a href="#">August 13, 2021</a>    | CDC recommends that people with advanced HIV disease or unsuppressed HIV RNA and others with “severe immune suppression” conditions receive a third dose after 2 mRNA doses no less than 28 days after the second dose. There was not enough evidence to recommend additional doses for those vaccinated with J&J. <ul style="list-style-type: none"> <li>Because our study participants were all fully vaccinated by June 30, 2021, our participants were ≥59 days since 2<sup>nd</sup> mRNA dose on August 13, 2021</li> </ul>  |
| <a href="#">September 24, 2021</a> | CDC recommends Pfizer boosters for those ≥6 months after completing the Pfizer primary series among: <ul style="list-style-type: none"> <li>≥65 years of age and residents in long-term care settings</li> <li>50-64 years of age with underlying medical conditions</li> </ul> CDC noted people 18-49 years with underlying conditions and people 18-64 who are at increased risk of COVID-19 exposure and transmission because of occupational or institutional settings *may* receive a booster  |
| <a href="#">October 21, 2021</a>   | CDC recommends Pfizer or Moderna boosters for those ≥6 months after completing an mRNA primary series among: <ul style="list-style-type: none"> <li>≥65 years of age and residents in long-term care settings</li> <li>≥18 years of age who live in long-term care settings</li> <li>≥18 years of age who have underlying medical conditions</li> <li>≥18 years of age who live or work in high-risk settings</li> </ul> CDC recommended booster shots for those ≥18 years of age who received J&J ≥2 months ago.<br>CDC states individuals may choose their booster vaccine type (i.e., mix and match dosing for boosters) |
| <a href="#">November 19, 2021</a>  | CDC recommends boosters for all those ≥18 years of age who received an mRNA primary series ≥6 months ago  |
| <a href="#">November 29, 2021</a>  | CDC strengthens recommendation from Nov 19, 2021, and expands to recommend all those ≥18 years of age who received J&J ≥2 months ago  |

**eFigure 2.** Distribution of Time From Fully Vaccinated to Receipt of Another Dose of Vaccine, by HIV Status, Among 133 994 Patients



**eTable 3.** Characteristics of 33 029 PWH Who Did and Did Not Receive a Vaccine Dose After Their Primary Series

| Characteristic                          | Overall<br>N = 33,029 <sup>1</sup> | Primary series only<br>N = 14,644 <sup>1</sup> | Primary series + dose < 5<br>months from full vaccination<br>series<br>N = 1,582 <sup>1</sup> | Primary series + dose ≥ 5<br>months from full vaccination<br>series<br>N = 16,803 | p-value <sup>2</sup> |
|---|------------------------------------|--|---|---|----------------------|
| <b>Age (years)</b>                      |                                    |  |   |   | <0.001               |
| 18-24                                   | 88 (0.3%)                          | 62 (0.4%)                                      | 6 (0.4%)  | 20 (0.1%)   |                      |
| 25-34                                   | 1,614 (4.9%)                       | 1,097 (7.5%)                                   | 71 (4.5%)   | 446 (2.7%)  |                      |
| 35-44                                   | 3,425 (10.4%)                      | 1,920 (13.1%)                                  | 205 (13.0%)   | 1,300 (7.7%)  |                      |
| 45-54                                   | 5,766 (17.5%)                      | 2,759 (18.8%)                                  | 389 (24.6%)   | 2,618 (15.6%)   |                      |
| 55-64                                   | 10,998 (33.3%)                     | 4,646 (31.7%)                                  | 691 (43.7%)   | 5,661 (33.7%)   |                      |
| 65-74                                   | 8,575 (26.0%)                      | 3,185 (21.7%)                                  | 193 (12.2%)   | 5,197 (30.9%)   |                      |
| 75+                                     | 2,563 (7.8%)                       | 975 (6.7%)                                     | 27 (1.7%)   | 1,561 (9.3%)  |                      |
| <b>Sex</b>                              |                                    |  |   |   | <0.001               |
| Male                                    | 30,676 (92.9%)                     | 13,482 (92.1%)                                 | 1,444 (91.3%)   | 15,750 (93.7%)  |                      |
| Female                                  | 2,353 (7.1%)                       | 1,162 (7.9%)                                   | 138 (8.7%)  | 1,053 (6.3%)  |                      |
| <b>Ethnicity and Race</b>               |                                    |  |   |   | <0.001               |
| Hispanic                                | 4,104 (12.4%)                      | 1,782 (12.2%)                                  | 223 (14.1%)   | 2,099 (12.5%)   |                      |
| Non-Hispanic Asian                      | 1,034 (3.1%)                       | 409 (2.8%)                                     | 77 (4.9%)   | 548 (3.3%)  |                      |
| Non-Hispanic Black/<br>African American | 13,545 (41.0%)                     | 6,518 (44.5%)                                  | 505 (31.9%)   | 6,522 (38.8%)   |                      |
| Non-Hispanic White                      | 12,968 (39.3%)                     | 5,293 (36.1%)                                  | 701 (44.3%)   | 6,974 (41.5%)   |                      |
| Other                                   | 1,076 (3.3%)                       | 493 (3.4%)                                     | 60 (3.8%)   | 523 (3.1%)  |                      |
| Unknown                                 | 302 (0.9%)                         | 149 (1.0%)                                     | 16 (1.0%)   | 137 (0.8%)  |                      |
| <b>Month fully vaccinated</b>           |                                    |  |   |   | <0.001               |
| January 2021                            | 359 (1.1%)                         | 164 (1.1%)                                     | 3 (0.2%)  | 192 (1.1%)  |                      |
| February 2021                           | 3,801 (11.5%)                      | 1,641 (11.2%)                                  | 25 (1.6%)   | 2,135 (12.7%)   |                      |

| Characteristic   | Overall<br>N = 33,029 <sup>1</sup> | Primary series only<br>N = 14,644 <sup>1</sup> | Primary series + dose < 5<br>months from full vaccination<br>series<br>N = 1,582 <sup>1</sup> | Primary series + dose ≥ 5<br>months from full vaccination<br>series<br>N = 16,803 | p-value <sup>2</sup> |
|--|------------------------------------|--|---|---|----------------------|
| March 2021   | 8,702 (26.3%)                      | 3,173 (21.7%)                                  | 48 (3.0%)   | 5,481 (32.6%)   |                      |
| April 2021   | 12,306 (37.3%)                     | 5,001 (34.2%)                                  | 768 (48.5%)   | 6,537 (38.9%)   |                      |
| May 2021   | 5,793 (17.5%)                      | 3,151 (21.5%)                                  | 604 (38.2%)   | 2,038 (12.1%)   |                      |
| June 2021  | 2,068 (6.3%)                       | 1,514 (10.3%)                                  | 134 (8.5%)  | 420 (2.5%)  |                      |
| <b>Primary vaccination series</b>                                |                                    |  |   |   | <0.001               |
| Pfizer   | 17,389 (52.6%)                     | 7,296 (49.8%)                                  | 885 (55.9%)   | 9,208 (54.8%)   |                      |
| Moderna  | 13,629 (41.3%)                     | 6,080 (41.5%)                                  | 660 (41.7%)   | 6,889 (41.0%)   |                      |
| J&J  | 2,011 (6.1%)                       | 1,268 (8.7%)                                   | 37 (2.3%)   | 706 (4.2%)  |                      |
| <b>COVID prior to fully vaccinated</b>                           | 2,272 (6.9%)                       | 1,086 (7.4%)                                   | 120 (7.6%)  | 1,066 (6.3%)  | <0.001               |
| <b>CD4 at ART initiation</b>                                     | 368.00 (202.00, 584.00)            | 373.00 (207.00, 594.00)                        | 356.00 (182.00, 555.00)   | 364.00 (198.00, 580.00)   | <0.001               |
| Unknown  | 14,910 (45.1%)                     | 6,275 (42.9%)                                  | 775 (49.0%)   | 7,860 (46.8%)   |                      |
| <b>AIDS before fully vaccinated</b>                              | 8,335 (25.2%)                      | 3,543 (24.2%)                                  | 343 (21.7%)   | 4,449 (26.5%)   | 0.008                |
| <b>CD4 at fully vaccinated</b>                                   | 636 (449, 858)                     | 634 (443, 858)                                 | 632 (430, 849)  | 640 (455, 859)  | 0.070                |
| Unknown  | 6,393 (19.3%)                      | 3,244 (22.2%)                                  | 204 (12.9%)   | 2,945 (17.5%)   |                      |
| <b>Suppressed (&lt;50 copies/mL) HIV RNA at fully vaccinated</b> | 26,052 (90.5%)                     | 10,810 (88.8%)                                 | 1,345 (90.5%)   | 13,897 (91.9%)  | <0.001               |
| Unknown  | 4,240 (12.8%)                      | 2,466 (16.8%)                                  | 95 (6.0%)   | 1,679 (10.0%)   |                      |
| <sup>1</sup> n (%); Median (IQR)                                 |                                    |  |   |   |                      |
| <sup>2</sup> Pearson's Chi-squared test; Wilcoxon rank sum test  |                                    |  |   |   |                      |

**eTable 4.** Breakthrough Infection Incidence Rates and 95% CIs, by Vaccine Type and HIV Status

|                          |         | # of breakthroughs | # of person-years | IR (95% CI) per 1,000 person years |
|--------------------------|---------|--------------------|-------------------|------------------------------------|
| <b>Moderna</b>           |         |                    |                   |                                    |
|                          | Overall | 1,140              | 33,262.7          | 34.3 (32.3, 36.3)                  |
|                          | PWH     | 369                | 9,391.2           | 39.3 (35.4, 43.5)                  |
|                          | PWoH    | 771                | 23,871.5          | 32.3 (30.1, 34.7)                  |
| <b>Pfizer</b>            |         |                    |                   |                                    |
|                          | Overall | 2,158              | 40,017.5          | 53.9 (51.7, 56.3)                  |
|                          | PWH     | 770                | 11,909.0          | 64.7 (60.2, 69.4)                  |
|                          | PWoH    | 1,388              | 28,108.5          | 49.4 (46.8, 52.1)                  |
| <b>J&amp;J</b>           |         |                    |                   |                                    |
|                          | Overall | 351                | 4999.5            | 70.2 (63.1, 78.0)                  |
|                          | PWH     | 102                | 1,327.9           | 76.8 (62.6, 93.3)                  |
|                          | PWoH    | 249                | 3,671.6           | 67.8 (59.7, 76.8)                  |
| <b>By calendar month</b> |         |                    |                   |                                    |
| Jan                      | Overall | 1                  | 25.0              | (1.0, 222.6)                       |
|                          | PWH     | 0                  | 7.5               | 0.0 (0.0, 492.3)                   |
|                          | PWoH    | 1                  | 17.5              | 57.0 (1.4, 317.8)                  |
| Feb                      | Overall | 13                 | 445.7             | 29.2 (15.5, 49.9)                  |
|                          | PWH     | 6                  | 139.5             | 43.0 (15.8, 93.6)                  |
|                          | PWoH    | 7                  | 306.2             | 22.9 (9.2, 47.1)                   |
| Mar                      | Overall | 52                 | 2322.9            | 22.4 (16.7, 29.4)                  |
|                          | PWH     | 17                 | 704.3             | 24.1 (14.1, 38.6)                  |
|                          | PWoH    | 35                 | 1618.6            | 21.6 (15.1, 30.1)                  |
| Apr                      | Overall | 107                | 5426.0            | 20.4 (16.7, 24.6)                  |
|                          | PWH     | 40                 | 1551.1            | 25.8 (18.4, 35.1)                  |
|                          | PWoH    | 67                 | 3694.9            | 18.1 (14.1, 23.0)                  |
| May                      | Overall | 95                 | 8338.5            | 11.4 (9.2, 139)                    |
|                          | PWH     | 28                 | 2418.1            | 11.6 (7.7, 16.7)                   |
|                          | PWoH    | 67                 | 5920.4            | 11.3 (8.8, 14.4)                   |
| June                     | Overall | 113                | 9082.6            | 12.4 (10.2, 15.0)                  |
|                          | PWH     | 43                 | 2624.6            | 16.4 (11.9, 22.1)                  |
|                          | PWoH    | 70                 | 6458.0            | 10.8 (8.4, 13.7)                   |
| Jul                      | Overall | 296                | 9557.6            | 72.1 (66.8, 77.8)                  |
|                          | PWH     | 136                | 2764.2            | 49.2 (41.3, 58.2)                  |
|                          | PWoH    | 160                | 6793.5            | 23.6 (20.0, 27.5)                  |
|                          | Overall | 684                | 9481.8            | 72.1 (66.8, 77.8)                  |



|      |         |       |        |                      |
|------|---------|-------|--------|----------------------|
| Aug  | PWH     | 259   | 2734.7 | 94.7 (83.5, 107.0)   |
|      | PWoH    | 425   | 6747.1 | 63.0 (57.1, 69.3)    |
| Sept | Overall | 497   | 9096.7 | 54.6 (49.9, 59.7)    |
|      | PWH     | 162   | 2621.2 | 61.8 (52.6, 72.1)    |
|      | PWoH    | 335   | 6475.5 | 51.7 (46.3, 57.6)    |
| Oct  | Overall | 294   | 9320.8 | 31.5 (28.0, 35.4)    |
|      | PWH     | 82    | 2683.4 | 30.6 (24.3, 37.9)    |
|      | PWoH    | 212   | 6637.3 | 31.9 (27.8, 36.5)    |
| Nov  | Overall | 232   | 8590.3 | 27.0 (23.6, 30.7)    |
|      | PWH     | 81    | 2463.1 | 32.9 (26.1, 40.9)    |
|      | PWoH    | 151   | 6127.2 | 24.6 (20.9, 28.9)    |
| Dec  | Overall | 1,265 | 7087.5 | 178.5 (168.8, 188.6) |
|      | PWH     | 387   | 2008.0 | 192.7 (174.0, 212.9) |
|      | PWoH    | 878   | 5079.6 | 172.8 (161.6, 184.7) |